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In the claims

This listing of claims will replace all prior versions, and listings, of claims in the application.

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Upon entry of the present amendment, the claims will stand as follows:

Please amend claims 19, 24 and 41 as follows:

Claims 1-18 (Cancelled)

19. (Currently Amended) A method for identifying a bioactivity or biomolecule of interest using high throughput screening of DNA comprising:

a) contacting a bioactive substrate that is fluorescent in the presence of the bioactivity or biomolecule of interest with a library containing a plurality of clones containing naturally occurring DNA from more than one organism, wherein each clone contains DNA from a single organism;

b) screening the library with a fluorescent analyzer that detects bioactive fluorescence; and

c) identifying clones detected as positive for bioactive fluorescence, wherein fluorescence is indicative of <u>a</u> naturally occurring DNA that encodes a bioactivity or biomolecule of interest.

20. (Previously Presented) The method of claim 19, further comprising obtaining DNA from a clone that is positive for an enzymatic activity of interest.

21. (Previously Presented) The method of claim 20, wherein the enzymatic activity of interest is from an enzyme selected from the group consisting of lipases, esterases, proteases, glycosidases, glycosyl transferases, phosphatases, kinases, diarylpropane peroxidases, epoxide hydrolases, nitrile hydratases, nitrilases, transaminases, amidases, and acylases.

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22. (Previously Presented) The method of claim 19, wherein the library is generated in a

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prokaryotic cell.

23. (Previously Presented) The method of claim 19, wherein the library contains at least

about 2 x 10⁶ clones.

24. (Currently Amended) The method of claim [[19]] 22, wherein the prokaryotic cell is

gram negative.

25. (Previously Presented) The method of claim 19, wherein the clones are encapsulated in a

gel microdrop.

26. (Previously Presented) The method of claim 19, wherein the analyzer screens up to about

15 million clones per hour.

27. (Previously Presented) The method of claim 19, wherein the clones are extremophiles.

28. (Previously Presented) The method of claim 27, wherein the extremophiles are

thermophiles.

29. (Previously Presented) The method of claim 27, wherein the extremophiles are

hyperthermophiles, psychrophiles, halophiles, psychrotrops, alkalophiles, or acidophiles.

30. (Previously Presented) The method of claim 19, wherein the bioactive substrate

comprises staining reagent C12-fluorescein-di-D-galactopyranoside (C12FDG).

31. (Previously Presented) The method of claim 19, wherein the bioactive substrate

comprises a lipophilic tail.

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32. (Previously Presented) The method of claim 19, wherein the clones and substrates are heated to enhance contacting of the substrate with the clones.

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- 33. (Previously Presented) The method of claim 32, wherein the heating is to a temperature of about 70°C.
- 34. (Previously Presented) The method of claim 32, wherein the heating is for about 30 minutes.
- 35. (Previously Presented) The method of claim 19, wherein the fluorescent analyzer comprises a fluorescence activated cell sorting (FACS) apparatus.
- 36. (Previously Presented) The method of claim 20, wherein the enzymatic activity of interest encoded by the DNA is stable at a temperature of at least about 60°C.
- 37. (Previously Presented) The method of claim 19, wherein the library is an expression library.
- 38. (Previously Presented) The method of claim 20, wherein the enzymatic activity of interest encoded by the DNA possesses enhanced enzymatic activity of interest compared to that of a wild-type enzyme.
- 39. (Previously Presented) The method of claim 19, wherein the method further comprises biopanning the expression library prior to contacting with the substrate.
- 40. (Previously Presented) The method of claim 19 further comprising obtaining DNA from a clone identified in step c) that is positive for an enzymatic activity of interest and comparing the enzymatic activity of a DNA expression product from the clone with that

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obtained from such a clone into whose DNA at least one nucleotide mutation has been

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introduced, wherein a difference in enzymatic activity is indicative of the effect upon the

enzymatic activity of interest caused by introduction of the at least one nucleotide

mutation.

41. (Currently Amended) The method of claim [[19]] 40, wherein the bioactivity encoded by

the DNA possesses the bioactivity of interest at a temperature at least 10 °C below the

temperature of optimal activity of the bioactivity encoded by the wild-type DNA.

42. (Cancelled).

43. (Previously Presented) The method of claim 19, wherein the library is a multispecies

library.

44. (Previously Presented) The method of claim 43, wherein the library is generated from a

mixed population of uncultured organisms.

45. (Previously Presented) The method of claim 43, wherein the library is generated from

isolates.

46. (Previously Presented) The method of claim 40, wherein the mutation is introduced by

error-prone PCR, oligonucleotide directed mutagenesis, assembly PCR, sexual PCR

mutagenesis, in vivo mutagenesis, cassette mutagenesis, recursive ensemble mutagenesis

and exponential ensemble mutagenesis.

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